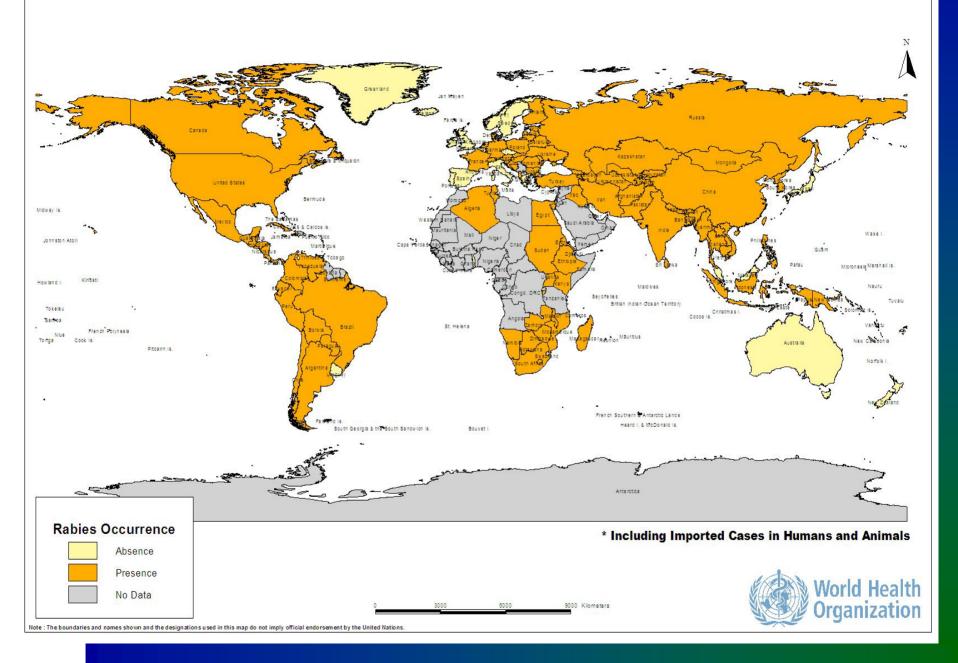
Rabies epidemiology, disease burden and vaccine utilization in Asia

Mary Elizabeth Miranda, DVM Philippines

Presence / Absence of Rabies* Worldwide - 2003



RABIES in ASIA

Over 30,000 die every year

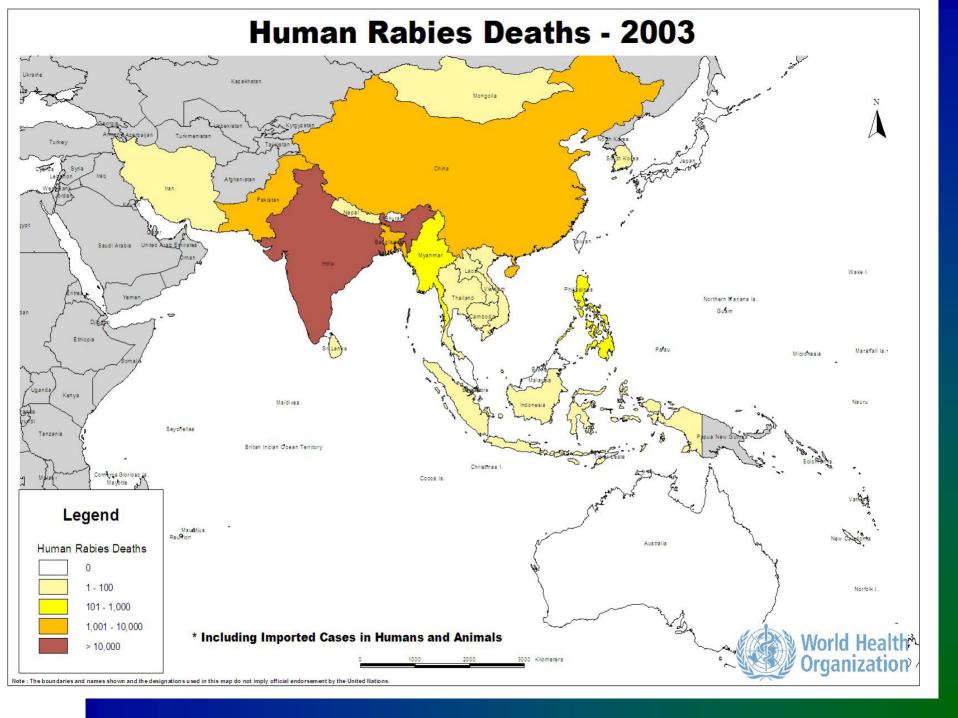


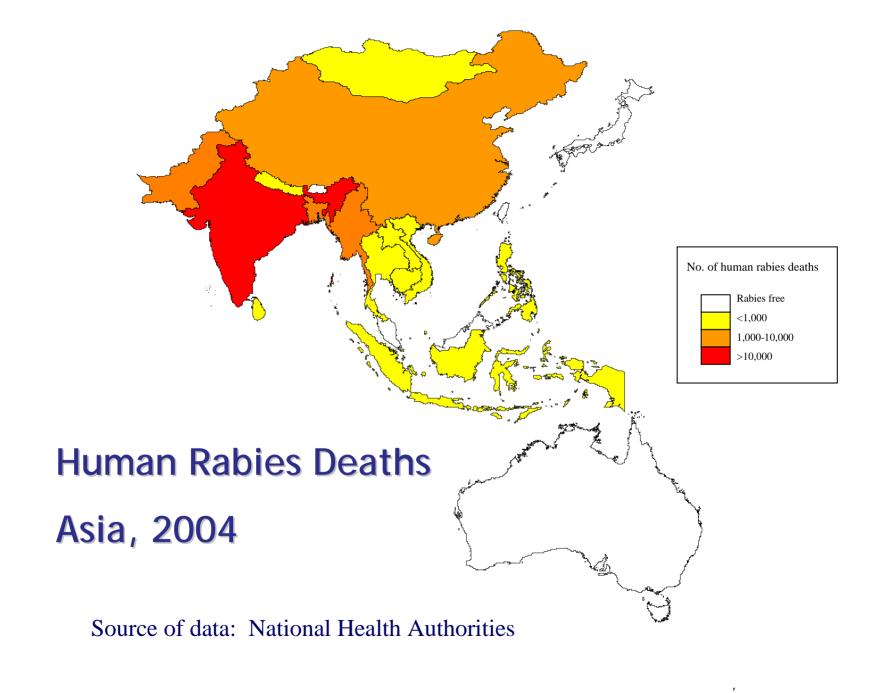
RABIES in ASIA

One Asian dies every 15 minutes, 50% likely to be a child under 15 years



San Lazaro Hospital, Department of Health, PHILIPPINES







More than 3 billion people in developing countries in Asia potentially exposed to dog rabies

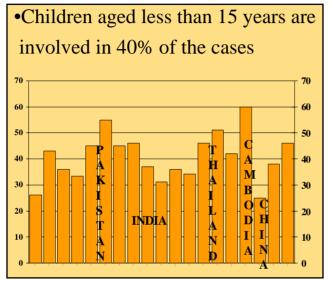
Rabies exposures

98% of deaths come from bites of rabid dog;

mostly children who come face to face with them



Rabies: a disease of childhood







Rabies a disease of neglected populations (rural & poor communities)

Human Rabies and Treatments, Asia, 2004

Country	Deaths	Rate/million	Post-exposure Treatment	Rate/mil
India	17,000	16.7	1,800,000	1,768
Pakistan	2,490	17	69,000	469
Bangladesh	1,550	12	60,000	455
Myanmar	1,100	23	5,000	102
China	2009	1.6	5,000,000	3,959
Philippines	248	3.3	102,148	1,338
Indonesia	40	0.2	8,800	43
Sri Lanka	76	4	80,000	4,200
Thailand	26	0.41	200,000	3,178
Vietnam	30	0. 38	635,000	8,105
Nepal	44	2.17	25,000	1,085
Cambodia	2	0.80	12,000	1,071
Lao People s Democratic Republic	2	1.26	3,000	540
Mongolia	2	0.80	62	25
TOTAL	24,609	6.5	7,692,010	1,881

Source: WHO World Survey of Rabies (RABNET online - www.who.int/rabnet)

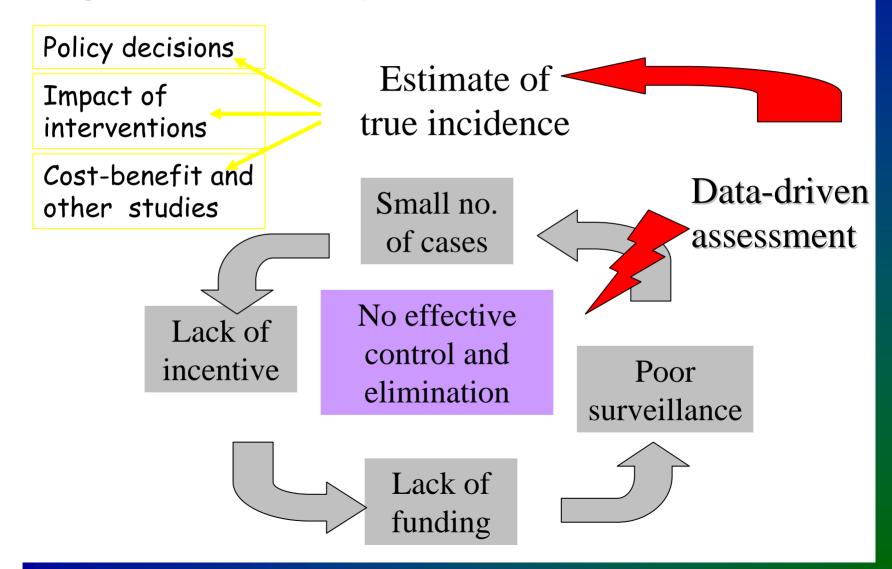
Post-exposure treatments in Asia

- Over 10M postexposure treatments annually
- 70 % of worldwide PET
- Majority still receive the nerve tissue derived rabies vaccine



800 post-exposure treatments every hour

Break "circle of neglect": ignorance-complacence



Need for a better rabies burden assessment:

WHO initiated work for the development of a global model for the re-assessment of rabies public health and economic burden in Africa and Asia.

We used the model developed by S. Cleaveland

Main parameters driving the model are

- Annual incidence of suspect rabid dog bites
- 'Rabies recognition probability'
- Probability of receiving successful post-exposure treatment
- Each parameter: a variable for which we have defined their is a minimum, a likely and a maximum value. Or a point probability
- For incidence of dog bites: from 10 to 250 per 100 000 most likely 100
- For P1: 39% to 64% most likely 50%
- For P10: 55% to 100% most likely 60 to 80%

Global outputs - deaths

Urban

Rural

- India = 1,058
- China = 1,324
- Other Asia = 853
- Africa = 5,886

- o India = 18,201
- o China = 1,257
- o Other Asia = 8,135
- o Africa = 17, 937

Urban:

9000: 16%

Rural:

46000: 84%

Global outputs:

Human Deaths: 55 000

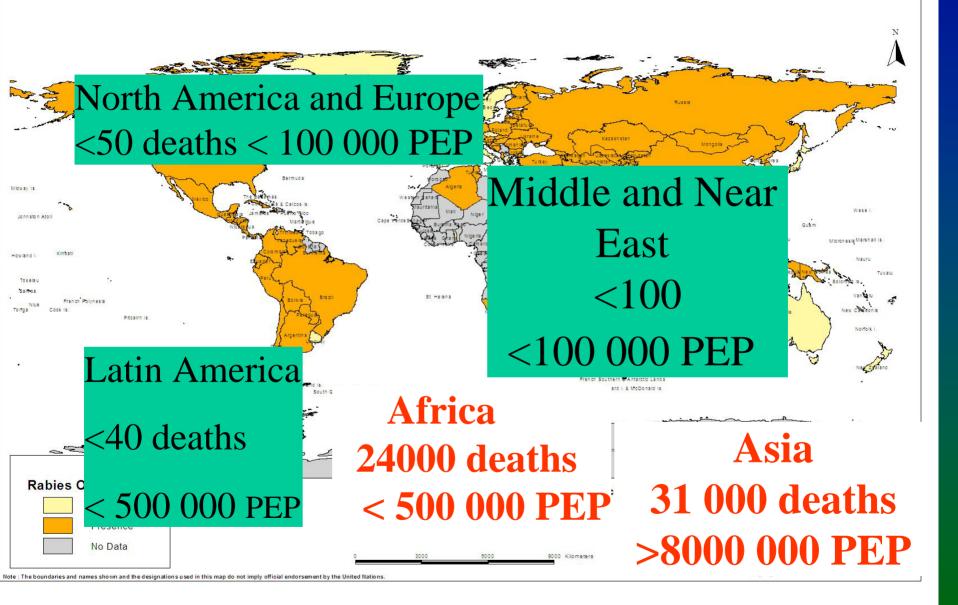
Total Asia: 31 000 (56% of total)

9 over 10 deaths in rural areas

Total Africa: 24 000 (44% of total)

3 over 4 deaths in rural areas

Presence / Absence of Rabies* Worldwide - 2003



Global outputs:

Cost burden: \$580 millions

Total Asia: 560 (96,5%)

Total Africa: 20 (3.5%)

DALY* scores

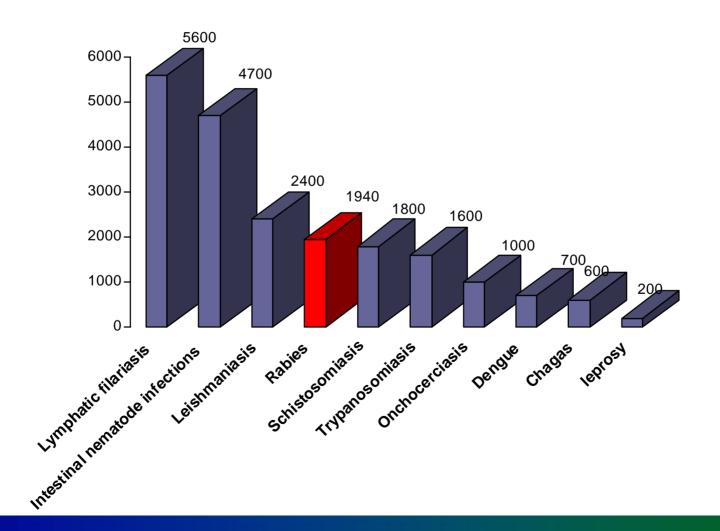
•DALY score (rabies death):

1,73 million -Africa:750,175, Asia: 975,796-(95% CI 408.000 - 3.360.000)

DALY score (fear factor and NTV side effects):

0,21 million (Asia: 0,178)

Ranking "Neglected Disease" burdens (DALYs x 1000)



Health burden without interventions

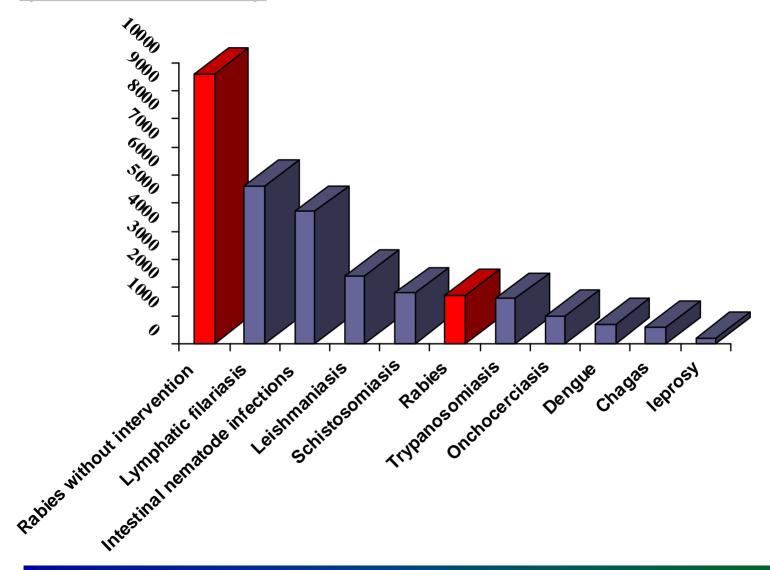
Total predicted human rabies deaths without human PET: 330,304

(95% CI: 141 000 - 565 000)

DALY score without intervention: 9.6 million

(95 %CI: 4.091.625 -16.370.517)

Ranking "Neglected Disease" burdens (DALYs x 1000)



Intersectoral National Rabies Control Programs in Asia

without

Bangladesh

Bhutan

Cambodia

India

Indonesia

Mongolia

Myanmar

with

- ✓ China
- ✓ Lao PDR
- ✓ Pakistan
- ✓ Philippines
- ✓ Nepal
- ✓ Sri Lanka
- ✓ Thailand
- ✓ Viet Nam

Common Issues and Concerns Rabies in Asia

- Not a priority disease
- Insufficient surveillance systems
- Limited access to modern rabies vaccines and supply problems
- Inadequate resources due to lack of political support
- Lack of consensus on strategy
- Weak intersectoral coordination
- Lack of public awareness and cooperation

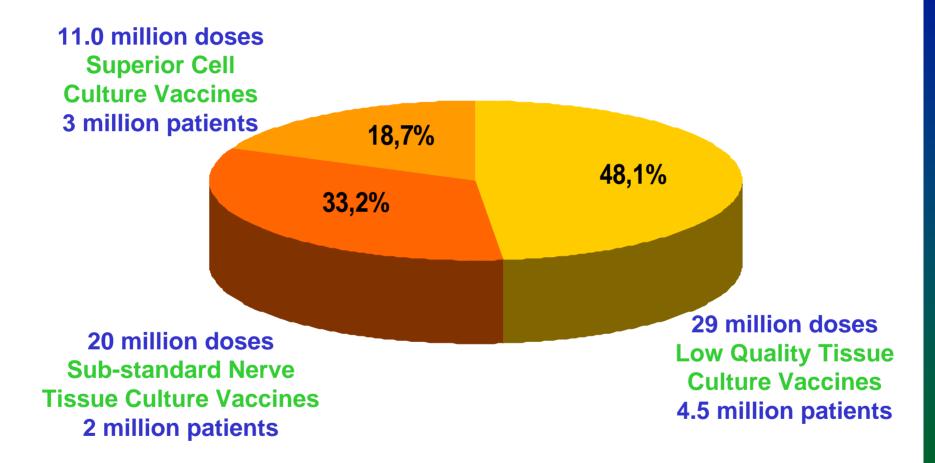
Nervous tissue derived vaccines still used in Asia in 2005

- O Bhutan
- ✓ Cambodia
- O China
- O India
- O Indonesia
- O Lao PDR
- ✓ Myanmar
- ✓ Nepal (phase out by Dec06)
- O Philippines
- O Sri Lanka
- O Thailand
- ✓ Viet Nam





Low quality vaccines are still used widely



Low quality and sub-standard nerve tissue vaccines dominate the global market. Less than 20% are superior cell culture vaccines (CCV).

Adapted from: Haupt W. In Rethinking Rabies Risks. Ed: Basgoz N. 2000

Rabies Control in Asia

Opportunities

- Utilize models of successful rabies elimination programs (Japan, Taiwan, Malaysia)
- Utilize current safe and efficacious vaccines
- Increase the number of patients receiving postexposure treatment
- Mobilize the support of civil society, private sector and government-sponsored initiatives
- Secure existing expertise on rabies/zoonoses

Summary of WHO Expert Consultation on Rabies WHO, Geneva, October 2004

published as

WHO Technical Report Series No.931

World Health Organization



 Report of this Expert Consultation supersedes the report of the 8th Expert Committee on Rabies held in 1992.

 Recommendations of this Consultation update and replace those contained in the 8th Report of the ECR

On vaccines the Consultation recommended that

- Production and use of all nerve tissue vaccines should be discontinued
- Only cell culture and purified embryonated eggs vaccines should be used in humans.
- Veterinary cell culture rabies vaccines should replace all nerve-tissue vaccines

Vaccines for human use:

- Rabies vaccines for human use should meet WHO requirements for the production and control of such vaccines, as well as the requirements that apply to the production of rabies vaccines as published in WHO Technical Report Series.
- NIH remains reference potency test for batch release.
 Minimum: 2,5 IU/single intramuscular immunizing dose.
- Candidate vaccines should demonstrate their safety, immunogenicity and or efficacy for human use in appropriate clinical trials using a vaccine of known efficacy as control and same WHO recommended PEP regimens.

<u>Immunoglobulin</u>

- Most of the new ERIGs are potent, highly purified, safe and considerably less expensive than HRIG
- However, a skin test should be conducted prior to administration of ERIG and F(ab´)2 products according to the guidelines of the manufacturer

Post-exposure prophylaxis

- All humans potentially exposed to rabies should receive prompt and thorough wound cleansing and antisepsis.
- This should be followed by passive immunization and a complete vaccine series using a potent and effective vaccine that meets WHO criteria applied using WHO regimens (2 IM, 2 IDs)
- Strict adherence to the WHO-recommended guidelines for optimal post-exposure rabies prophylaxis virtually guarantees protection from the disease.

Post-exposure prophylaxis

3 Categories of contact:

No exposure/category I: no treatment

Minor exposure/category II: vaccine only

Severe exposure/ category III: vaccine and

RIG

Bat contact under category III

Intradermal immunization for Injecting vaccine intrademally against resistance pre and post-exposure prophylaxis



 Unique ID dose for WHO recommended multidose intradermal Pre and Post EP regimens is 0,1 mL per id site

 Intradermal Post EP only with WHO recommended vaccines (PCECV Chiron, PVRV Aventis, HDCV from both companies).

Recommendations for safe clinical management of rabies patients

- The care of humans diagnosed with rabies often creates great anxiety in a hospital setting, involving not only medical and nursing staff but the media and the public.
- Pre-exposure immunization against rabies of nursing staff and health-care personnel in hospitals may be provided to those who, after careful investigation, are considered most at risk.
- However the need to strictly adhere to proper barrier nursing methods for patient care, should be emphasized as equally if not more important in caring for rabies patients.

Postmortem management of bodies of patients who have died of rabies

- When performing necropsies wearing protective clothing, goggles, a face mask and thick gloves should provide sufficient protection. Instruments must be autoclaved or boiled after use.
- Early disposal of the body by cremation or burial is recommended. Embalming should be discouraged.

Dog rabies control the Consultation:



 Stated that mass immunisation: the most important single measure

 Recommended the use as a complementary measure of oral vaccines in dogs

Dog population management the Consultation:

 Acknowledged the increasing number of animal birth control programmes (capture, neuter, vaccinate, release) and the encouraging results obtained

 Reiterated that indiscriminate mass dog culling is ineffective for dog rabies control

Too many needlessly die of rabies in Asia



EVERY ONE OF THESE DEATHS CAN BE PREVENTED